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## (54) Title: NOVEL INHIBITORS OF UBIQUITIN ISOPEPTIDASES

(57) Abstract: A novel class of inhibitors of ubiquitin isopeptidases is disclosed that cause tumor cell death via molecular mechanisms independent of p53. Specifically, compounds containing an a, 8-unsaturated ketone with a sterically accessible electrophilic B-carbon and related compounds are identified herein. Pharmaceutical compositions comprising the inhibitor compounds and methods of using the compounds for treating a variety of disease states are disclosed.